Amendments to the Claims

Claims 1-7 and 17-47 were previously cancelled. With this amendment, please amend claims 8, 48, 55, and 60, 70, 71, and 73-75; and cancel claim 10, as indicated below:

Claims 1-7. Cancelled.

Claim 8. (currently amended) A method of directing differentiation of human embryonic cells to a specific cell type, comprising:

- a. obtaining a hES cell line from inner cell mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell line;
- c. aggregating the hES cells in suspension, without adherence to the culture vessel, to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies;
 - d. dissociating the embryoid bodies to provide dissociated embryonic cells;
 - e. culturing said dissociated embryonic cells as a monolayer; and
- f. exposing said monolayer to at least one exogenous factor for an effective period of time to direct differentiation of said dissociated embryonic cells to form the specific cell type comprising a marker for terminally differentiated cells of the specific cell type.

Claim 9. (original) A method according to claim 8, wherein the embryoid bodies are formed in a suspension culture.

Claim 10. Cancelled.

Claim 11. (original) A method according to claim 8, wherein the exogenous factor is a growth factor.

Claim 12. (original) A method according to claim 8, wherein the exogenous factor is an interleukin.

Claim 13. (original) A method according to claim 11, wherein the exogenous factor is nerve growth factor.

Claim 14. (original) A method according to claim 8, wherein the exogenous factor is retinoic acid.

Claim 15. (original) A method according to claim 8, wherein the differentiated cells are neuronal cell type.

Claim 16. (original) A method according to claim 15, wherein the differentiated cells have neuronal processes.

Claims 17-47. Cancelled.

Claim 48. (currently amended) A method of directing differentiation of human embryonic cells to human ectoderm cells, comprising:

- a. obtaining a hES cell line from inner cell mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell line;
- c. aggregating the hES cells in suspension, without adherence to the culture vessel, to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies;
 - d. dissociating the embryoid bodies to provide dissociated embryonic cells;
 - e. culturing said dissociated embryonic cells as a monolayer; and
- f. exposing said monolayer to at least one exogenous factor for an effective period of time to direct differentiation of said dissociated embryonic cells to form human ectoderm cells comprising a marker for terminally differentiated human ectoderm cells.

Claim 49. (withdrawn) A method according to claim 48, wherein, in causing, said embryonic cells form human epidermal skin cells.

Claim 50. (currently amended) A method according to claim 49, wherein, in exposing, the at least one exogenous factor includes EGF.

Claim 51. (previously presented) A method according to claim 48, wherein, in causing, said embryonic cells form human brain cells.

Claim 52. (previously presented) A method according to claim 51, wherein, in exposing, the at least one exogenous factor includes at least one of RA and NGF.

Claim 53. (withdrawn) A method according to claim 48, wherein, in causing, said embryonic cells form human adrenal cells.

Claim 54. (previously presented) A method according to claim 53, wherein, in exposing, the at least one exogenous factor includes RA.

Claim 55. (currently amended) A method of directing differentiation of human embryonic cells to human endoderm cells, comprising:

- a. obtaining a hES cell line from inner cell mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell line:
- c. aggregating the hES cells in suspension, without adherence to the culture vessel, to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies;
 - d. dissociating the embryoid bodies to provide dissociated embryonic cells;
 - e. culturing said dissociated embryonic cells as a monolayer; and
- f. exposing said monolayer to at least one exogenous factor for an effective period of time to direct differentiation of said dissociated embryonic cells to form human endoderm cells comprising a marker for terminally differentiated human endoderm cells.

- Claim 56. (withdrawn) A method according to claim 55, wherein, in causing, said embryonic cells form human liver cells.
- Claim 57. (previously presented) A method according to claim 56, wherein, in exposing, the at least one exogenous factor includes at least one of HGF and NGF.
- Claim 58. (withdrawn) A method according to claim 55, wherein, in causing, said embryonic cells form human pancreatic cells.
- Claim 59. (previously presented) A method according to claim 58, wherein, in exposing, the at least one exogenous factor includes at least one of HGF and NGF.
- Claim 60. (currently amended) A method of directing differentiation of human embryonic cells to human mesoderm cells, comprising:
 - a. obtaining a hES cell line from inner cell mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell
 line;
- c. aggregating the hES cells in suspension, without adherence to the culture vessel, to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies;
 - d. dissociating the embryoid bodies to provide dissociated embryonic cells;
 - e. culturing said dissociated embryonic cells as a monolayer; and
 - f. exposing said monolayer to at least one exogenous factor for an effective period of time to direct differentiation of said dissociated embryonic cells to form human mesoderm cells comprising a marker for terminally differentiated human mesoderm cells.
- Claim 61. (withdrawn) A method according to claim 60, wherein, in causing, said embryonic cells form human chondrocytes.

Claim 62. (previously presented) A method according to claim 61, wherein, in exposing, the at least one exogenous factor includes BMP-4.

Claim 63. (withdrawn) A method according to claim 60, wherein, in causing, said embryonic cells form human kidney cells.

Claim 64. (withdrawn) A method according to claim 60, wherein, in causing, said embryonic cells form human Mullerian duct cells.

Claim 65. (previously presented) A method according to claim 60, wherein, in causing, said embryonic cells form human blood cells.

Claim 66. (withdrawn) A method according to claim 60, wherein, in causing, said embryonic cells form human heart muscle cells.

Claim 67. (previously presented) A method according to claim 66, wherein, in exposing, the at least one exogenous factor includes at least one of TGF- β and activin-A.

Claim 68. (withdrawn) A method according to claim 60, wherein, in causing, said embryonic cells form human skeletal muscle cells.

Claim 69. (previously presented) A method according to claim 68, wherein, in exposing, the at least one exogenous factor includes at least one of TGF-β and activin-A.

70. (currently amended) A method of directing differentiation of human embryonic cells to human neuronal cells comprising:

- a. obtaining a hES cell line from inner cell mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell line;

- c. aggregating the hES cells in suspension, without adherence to the culture vessel, to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies;
 - dissociating the embryoid bodies to provide dissociated embryonic cells;
 - e. culturing said dissociated embryonic cells as a monolayer; and
- f. exposing said monolayer to at least one exogenous factor for an effective period of time to direct differentiation of said dissociated embryonic cells to form the neuronal cells.
- 71. (currently) A method of directing differentiation of human embryonic cells to human muscle cells comprising:
 - a. obtaining a hES cell line from inner cell mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell line;
- c. aggregating the hES cells in suspension, without adherence to the culture vessel, to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies;
 - d. dissociating the embryoid bodies to provide dissociated embryonic cells;
 - e. culturing said dissociated embryonic cells as a monolayer; and
- f. exposing said monolayer to at least one exogenous factor for an effective period of time to direct differentiation of said dissociated embryonic cells to form the muscle cells.
- 72. (previously presented) A method according to claim 71, wherein the muscle cells are cardiomyocytes.
- 73. (currently amended) A method of directing differentiation of human embryonic cells to human pancreatic cells comprising:
 - a. obtaining a hES cell line from inner <u>cell</u> mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell line;

- c. aggregating the hES cells in suspension, without adherence to the culture vessel, to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies;
 - dissociating the embryoid bodies to provide dissociated embryonic cells;
 - e. culturing said dissociated embryonic cells as a monolayer; and
- f. exposing said monolayer to at least one exogenous factor for an effective period of time to direct differentiation of said dissociated embryonic cells to form the pancreatic cells.
- 74. (currently amended) A method of making human embryonic bodies from human embryonic stem cells comprising:
 - a. obtaining a hES cell line from inner cell mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell line:
- c. aggregating the hES cells in suspension, without adherence to the culture vessel, so as to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies.
- 75. (currently amended) A method of making human embryonic cells from human embryonic bodies comprising:
 - a. obtaining a hES cell line from inner cell mass-stage human embryos;
- b. chemically dissociating the human embryonic stem cells from said cell line;
- c. aggregating the hES cells in suspension, without adherence to the culture vessel, to form embryoid bodies and initial differentiation of the aggregated hES in the embryoid bodies;
- d. dissociating the embryoid bodies to provide dissociated embryonic cells;
 and
 - e. culturing said dissociated embryonic cells.